Schulwitz U.S. DEPARTMENT OF COMMERCE Patent and Trademark Office SEARCH REQUEST FORM Reducator Number: 99/047802 Art Unit: 16/ (/ 4/12/04 X 111 4 C70 Inv. Kobert Show Please write a detailed statement of search topic. Describe specifically as possible the subject matter to be searched. Define any terms that may have a special meaning. Give examples or relevent citations, authors, keywords, etc., if known. For sequences, please attach a copy of the sequence. You may include a copy of the broadest and/or most relevent claim(s). Please reach the uttached empount of formula T cel 11) to I treat cancer/neighborn/
(cl 11) to I treat cancer/neighborn/
metastasis of the methods of claim 19,20
netastasis of the methods of claim 19,20
21 Learch in capitus, medlini campoiner
21 Learch in capitus,

Rish leavel Approved

TK ParaSPE, 1615

STAFF USE ONLY

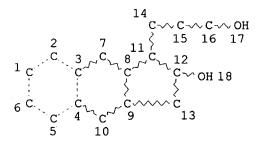
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Searcher:	STIC	<u> </u>
Terminal time:	CM-1	402,76 STN
Elapsed time:	Pre-S	Dialog
CPU time:	Type of Search	APS
Total time:	N.A. Sequence	Geninfo
Number of Scarches:	A.A. Sequence	- <u> </u>
Number of Databases:	Structure	DARC/Questel
	Bibliographic	Other

NODE ATTRIBUTES:
CONNECT IS X3 RC AT 16
DEFAULT MLEVEL IS ATOM
DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES: RING(S) ARE ISOLATED OR EMBEDDED NUMBER OF NODES IS 18

STEREO ATTRIBUTES: NONE

L3 87 SEA FILE=REGISTRY SSS FUL L1 L5 STR



NODE ATTRIBUTES:
CONNECT IS E2 RC AT 5
CONNECT IS X3 RC AT 16
DEFAULT MLEVEL IS ATOM
DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES: RING(S) ARE ISOLATED OR EMBEDDED NUMBER OF NODES IS 18

STEREO ATTRIBUTES: NONE

L6 2 SEA FILE=REGISTRY SUB=L3 SSS FUL L5

L7 ST

0 @25

VAR G2=O/N/S/CVAR G3=24/25NODE ATTRIBUTES: CONNECT IS X3 RC AT 16

CONNECT IS E2 RC AT 20 RC AT CONNECT IS E1 24 CONNECT IS E1 RC AT

DEFAULT MLEVEL IS ATOM

DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:

RING(S) ARE ISOLATED OR EMBEDDED NUMBER OF NODES IS 25

STEREO ATTRIBUTES: NONE

L836 SEA FILE=REGISTRY SUB=L3 SSS FUL L7

L12 12 SEA FILE=MEDLINE ABB=ON PLU=ON (L6 OR L8)

(L13 O SEA FILE MEDLINE ABBON PLUON L12 AND (?CANCER? OR ?NEOPLAS? OR ?CARCIN? OR ?TUMOR? OR ?TUMOUR? OR ?METAST?)

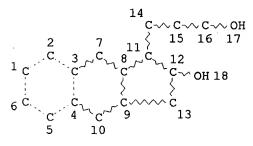
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NODE ATTRIBUTES: CONNECT IS X3 RC AT 16 DEFAULT MLEVEL IS ATOM DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES: RING(S) ARE ISOLATED OR EMBEDDED NUMBER OF NODES IS 18

STEREO ATTRIBUTES: NONE

87 SEA FILE=REGISTRY SSS FUL L1 L3 L5 STR



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GRAPH ATTRIBUTES: RING(S) ARE ISOLATED OR EMBEDDED NUMBER OF NODES IS 18

STEREO ATTRIBUTES: NONE

L6 2 SEA FILE=REGISTRY SUB=L3 SSS FUL L5 L7

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GRAPH ATTRIBUTES:
RING(S) ARE ISOLATED OR EMBEDDED
NUMBER OF NODES IS 25

STEREO ATTRIBUTES: NONE

L8

36 SEA FILE=REGISTRY SUB=L3 SSS FUL L7

L10

23 SEA FILE=HCAPLUS ABB=ON PLU=ON (L6 OR L8)(L)(BAC OR DMA OR

PAC OR PKT OR THU)/RL

/ L11

2 SEA FILE=HCAPLUS ABB=ON PLU=ON L10 AND (?CANCER? OR ?NEOPLAS?

OR ?CARCIN? OR ?TUMOR? OR ?TUMOUR? OR ?METAST?)

=> d 111 ibib ab hitind hitstr 1-2

L11 ANSWER 1 OF 2 HCAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER:

2003:678597 HCAPLUS

DOCUMENT NUMBER:

139:219309

TITLE:

Prostacyclin derivative-containing compositions and

methods of using the same for the treatment of

cancer

INVENTOR(S):

Shorr, Robert; Rothblatt, Martine United Therapeutics Corporation, USA

PATENT ASSIGNEE(S): SOURCE:

PCT Int. Appl., 65 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.

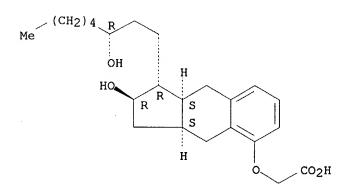
KIND DATE

APPLICATION NO. DATE

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             CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
             GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
             LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,
             PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT, TZ,
             UA, UG, UZ, VC, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU,
             TJ, TM
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, BG,
             CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC,
             NL, PT, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GO, GW.
             ML, MR, NE, SN, TD, TG
     US 2003166728
                            20030904
                      A1
                                           US 2002-47802
                                                            20020116
PRIORITY APPLN. INFO.:
                                        US 2002-47802
                                                        A 20020116
OTHER SOURCE(S):
                         MARPAT 139:219309
     The present invention is directed to a pharmaceutical compn. contg. a
     cancer-treating effective amt. of a class of prostacyclin derivs.,
     and a pharmaceutically acceptable carrier, and to kits and methods of
     employing the same for the treatment of cancer. For example,
     the prostacyclin deriv. inhibited protein degrdn. and promoted apoptosis
     of human amelanotic melanoma cells.
     ICM A61K
IC
     63-6 (Pharmaceuticals)
CC
     Section cross-reference(s): 1, 25
     prostacyclin protein degrdn inhibitor cancer
ST
IT
     Prostaglandins
     RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU
     (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES
        (I; compns. contg. prostacyclin deriv. for cancer treatment)
IT
     Melanoma
        (amelanotic; compns. contg. prostacyclin deriv. for cancer
        treatment)
IT
     Antitumor agents
     Human
      Neoplasm
        (compns. contg. prostacyclin deriv. for cancer treatment)
IT
     Collagens, biological studies
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (degrdn., inhibition of; compns. contg. prostacyclin deriv. for
        cancer treatment)
ΙT
     Drug delivery systems
        (inhalants; compns. contg. prostacyclin deriv. for cancer
IT
    Drug delivery systems
        (injections, i.v.; compns. contg. prostacyclin deriv. for
        cancer treatment)
IT
    Drug delivery systems
        (injections, s.c.; compns. contg. prostacyclin deriv. for
       cancer treatment)
IΤ
    Drug delivery systems
        (kits; compns. contg. prostacyclin deriv. for cancer
       treatment)
    Neoplasm
        (metastasis, inhibition of; compns. contg. prostacyclin
       deriv. for cancer treatment)
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IT Drug delivery systems (oral; compns. contq. prostacyclin deriv. for cancer treatment) IT Drug delivery systems (parenterals; compns. contg. prostacyclin deriv. for cancer treatment) IT Apoptosis (promotion of; compns. contg. prostacyclin deriv. for cancer treatment) TT Extracellular matrix (protein degrdn. in, inhibition of; compns. contg. prostacyclin deriv. for cancer treatment) IT 343247-13-2P RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (compns. contg. prostacyclin deriv. for cancer treatment) ΙT 823-96-1 6971-51-3, 3-Methoxybenzyl alcohol 22348-32-9 223734-62-1 RL: RCT (Reactant); RACT (Reactant or reagent) (compns. contg. prostacyclin deriv. for cancer treatment) ΙT 94956-98-6P 101692-01-7P 101692-02-8P 101692-03-9P 101758-87-6P 136911-16-5P 153974-48-2P 223734-55-2P 223734-56-3P 223734-57-4P 223734-58-5P 223734-59-6P 223734-60-9P 223734-61-0P RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent) (compns. contg. prostacyclin deriv. for cancer treatment) IT 343247-13-2P RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (compns. contg. prostacyclin deriv. for cancer treatment) RN 343247-13-2 HCAPLUS CN Acetic acid, [(1R,2R,3aS,9aS)-2,3,3a,4,9,9a-hexahydro-2-hydroxy-1-[(3R)-3hydroxyoctyl]-1H-benz[f]inden-5-yl]oxy]- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



L11 ANSWER 2 OF 2 HCAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER:

2002:720700 HCAPLUS

DOCUMENT NUMBER:

138:19862

TITLE:

The Prostacyclin Analogue Treprostinil Blocks NF.kappa.B Nuclear Translocation in Human Alveolar Macrophages

AUTHOR(S): Raychaudhuri, Baisakhi; Malur, Anagha; Bonfield,

Tracey L.; Abraham, Susamma; Schilz, Robert J.;

Farver, Carol F.; Kavuru, Mani S.; Arroliga, Alejandro

C.; Thomassen, Mary Jane

CORPORATE SOURCE: Department of Pulmonary and Critical Care Medicine,

The Cleveland Clinic Foundation, Cleveland, OH,

44195-5038, USA

426102 Journal of Biological Chemistry (2002), 277(36),

33344-33348

CODEN: JBCHA3; ISSN: 0021-9258

PUBLISHER:

American Society for Biochemistry and Molecular

Biology

DOCUMENT TYPE:

LANGUAGE:

SOURCE:

Journal English

Primary pulmonary hypertension (PPH) is characterized by increased pulmonary arterial pressure and vascular resistance. We and others have obsd. that inflammatory cytokines and infiltrates are present in the lung tissue, but the significance is uncertain. Treprostinil (TRE), a prostacyclin analog with extended half-life and chem. stability, has shown promise in the treatment of PPH. We hypothesize that TRE might exert beneficial effects in PPH by antagonizing inflammatory cytokine prodn. in the lung. Here we show that TRE dose-dependently inhibits inflammatory cytokine (tumor necrosis factor-.alpha., interleukin-1.beta., interleukin-6, and granulocyte macrophage colony-stimulating factor) secretion and gene expression by human alveolar macrophages. TRE blocks NF.kappa.B activation, but I.kappa.B-.alpha. phosphorylation and degrdn. are unaffected. Moreover, TRE does not affect the formation of the NF.kappa.B.cntdot.DNA complex but blocks nuclear translocation of p65. These results are the first to illustrate the anti-cytokine actions of TRE

in down-regulating NF.kappa.B, not through its inhibitory component or by direct binding but by blocking nuclear translocation. These data indicate that inflammatory mechanisms may be important in the pathogenesis of PPH

and cytokine antagonism by blocking NF.kappa.B may contribute to the efficacy of TRE therapy in PPH.

2-9 (Mammalian Hormones) CC Interleukin 1.beta.

Interleukin 6

Tumor necrosis factors

RL: BSU (Biological study, unclassified); BIOL (Biological study) (prostacyclin analog treprostinil blocks inflammatory cytokine secretion and NF.kappa.B nuclear translocation in human alveolar macrophages in relation to it use in primary pulmonary hypertension treatment)

81846-19-7, Treprostinil

RL: DMA (Drug mechanism of action); BIOL (Biological study) (prostacyclin analog treprostinil blocks inflammatory cytokine secretion and NF.kappa.B nuclear translocation in human alveolar macrophages in relation to it use in primary pulmonary hypertension treatment)

IT **81846-19-7**, Treprostinil

RL: DMA (Drug mechanism of action); BIOL (Biological study) (prostacyclin analog treprostinil blocks inflammatory cytokine secretion and NF.kappa.B nuclear translocation in human alveolar macrophages in relation to it use in primary pulmonary hypertension treatment)

81846-19-7 HCAPLUS RN

CN Acetic acid, [[(1R,2R,3aS,9aS)-2,3,3a,4,9,9a-hexahydro-2-hydroxy-1-[(3S)-3-hydroxyoctyl]-1H-benz[f]inden-5-yl]oxy]- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

REFERENCE COUNT:

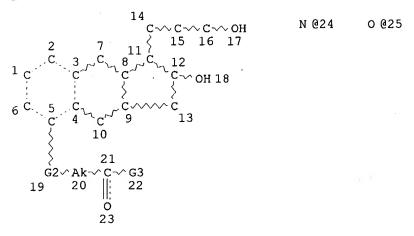
32 THERE ARE 32 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

NODE ATTRIBUTES: CONNECT IS X3 RC AT 16 DEFAULT MLEVEL IS ATOM DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES: RING(S) ARE ISOLATED OR EMBEDDED NUMBER OF NODES IS 18

STEREO ATTRIBUTES: NONE

L3 87 SEA FILE=REGISTRY SSS FUL L1 L7 STR



VAR G2=O/N/S/C
VAR G3=24/25
NODE ATTRIBUTES:
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CONNECT IS E2 RC AT 20
CONNECT IS E1 RC AT 24
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DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES: RING(S) ARE ISOLATED OR EMBEDDED NUMBER OF NODES IS 25 STEREO ATTRIBUTES: NONE

L8 36 SEA FILE=REGISTRY SUB=L3 SSS FUL L7

L17 85 SEA L8

L18 6 SEA L17 AND (CANCER OR ANTICANC? OR NEOPLAS? OR ANTINEOPLAS?
OR TUMOR? OR ANTITUM? OR METAST? OR CARCIN?)

=> dup rem 118

PROCESSING COMPLETED FOR L18

L19 6 DUP REM L18

6 DUP REM L18 (0 DUPLICATES REMOVED)
ANSWERS '1-2' FROM FILE BIOSIS
ANSWERS '3-6' FROM FILE USPATFULL

(=> d 119 bib ab 1-6)

- L19 ANSWER 1 OF 6 BIOSIS COPYRIGHT 2004 BIOLOGICAL ABSTRACTS INC. on STN
- AN 2003:417438 BIOSIS
- DN PREV200300417438
- TI 27th Meeting of the Oesterreichische Gesellschaft fuer Lungenerkrankungen und Tuberkulose, Alpbach, Tyrol, Austria, May, 29-June, 1, 2003.
- AU Oesterreichische Gesellschaft fuer Lungenerkrankungen und Tuberkulose
- SO Wiener Klinische Wochenschrift, (30 Mai 2003) Vol. 115, No. 10, pp. A IV-A XIII. print.

 Meeting Info: 27th Meeting of the Oesterreichische Cosellaghaft fuor

Meeting Info.: 27th Meeting of the Oesterreichische Gesellschaft fuer Lungenerkrankungen und Tuberkulose. Alpbach, Tyrol, Austria. May 29-June 01, 2003. Oesterreichische Gesellschaft fuer Lungenerkrankungen und Tuberkulose.

CODEN: WKWOAO. ISSN: 0043-5325.

- DT Conference; (Meeting)
 Conference; (Meeting Summary)
- LA English
- ED Entered STN: 10 Sep 2003

Last Updated on STN: 10 Sep 2003

- AB This meeting contains abstracts of 33 papers, written in German and English, on a variety of topics in lung diseases in the human patient, including asthma bronchiale, sleep apnea syndrome, bronchial carcinoma, pancreatic tumor, tuberculosis, lung cancer, respiratory failure, actinomycosis, HIV, pulmonary arterial hypertension, ventilation, autofluorescence bronchoscopy, bronchoscopy, endoscopy simulator, smoking cessation therapy, nebulizer therapy, bronchoalveolar lavage, mycophenolate mofetil, treprostinil, high-altitude medicine, and genetics.
- L19 ANSWER 2 OF 6 BIOSIS COPYRIGHT 2004 BIOLOGICAL ABSTRACTS INC. on STN
- AN 2002:535469 BIOSIS
- DN PREV200200535469
- TI The prostacyclin analogue treprostinil blocks NFkappaB nuclear translocation in human alveolar macrophages.
- AU Raychaudhuri, Baisakhi; Malur, Anagha; Bonfield, Tracey L.; Abraham, Susamma; Schilz, Robert J.; Farver, Carol F.; Kavuru, Mani S.; Arroliga, Alejandro C.; Thomassen, Mary Jane [Reprint author]
- CS Dept. of Pulmonary and Critical Care Medicine, Cleveland Clinic Foundation, 9500 Euclid Ave., Desk A90, Cleveland, OH, 44195-5038, USA thomasm@ccf.org
- SO Journal of Biological Chemistry, (September 6, 2002) Vol. 277, No. 36, pp. 33344-33348. print. CODEN: JBCHA3. ISSN: 0021-9258.

```
DT Article
LA English
```

ED Entered STN: 16 Oct 2002 Last Updated on STN: 16 Oct 2002

AB Primary pulmonary hypertension (PPH) is characterized by increased pulmonary arterial pressure and vascular resistance. We and others have observed that inflammatory cytokines and infiltrates are present in the lung tissue, but the significance is uncertain. Treprostinil (TRE), a prostacyclin analogue with extended half-life and chemical stability, has shown promise in the treatment of PPH. We hypothesize that TRE might exert beneficial effects in PPH by antagonizing inflammatory cytokine production in the lung. Here we show that TRE dose-dependently inhibits inflammatory cytokine (tumor necrosis factor-alpha, interleukin-1beta, interleukin-6, and granulocyte macrophage colony-stimulating factor) secretion and gene expression by human alveolar macrophages. TRE blocks NFkappaB activation, but IkappaB-alpha phosphorylation and degradation are unaffected. Moreover, TRE does not affect the formation of the NFkappaBcntdotDNA complex but blocks nuclear translocation of p65. These results are the first to illustrate the anti-cytokine actions of TRE in down-regulating NFkappaB, not through its inhibitory component or by direct binding but by blocking nuclear translocation. These data indicate that inflammatory mechanisms may be important in the pathogenesis of PPH and cytokine antagonism by blocking NFkappaB may contribute to the efficacy of TRE therapy in PPH.

```
L19 ANSWER 3 OF 6 USPATFULL on STN
```

AN 2003:307038 USPATFULL

TI Method of using prostacyclin to treat respiratory syncytial virus infections

IN Peebles, Ray Stokes, JR., Nashville, TN, UNITED STATES Hashimoto, Koichi, Fukushima, JAPAN Graham, Barney S., Rockville, MD, UNITED STATES

PI US 2003216474 A1 20031120 AI US 2003-389295 A1 20030314 (10)

PRAI US 2002-364395P 20020315 (60)

DT Utility

FS APPLICATION

LREP WADDEY & PATTERSON, 414 UNION STREET, SUITE 2020, BANK OF AMERICA PLAZA, NASHVILLE, TN, 37219

CLMN Number of Claims: 20 ECL Exemplary Claim: 1 DRWN 9 Drawing Page(s)

LN.CNT 1178

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The present invention discloses methods and a kit for treating a respiratory syncytial virus infection. The method comprises providing an infection modulator, and administering a therapeutically effective amount of the infection modulator, wherein the respiratory syncytial virus infection is suppressed or precluded. The kit for suppressing a respiratory syncytial virus infection comprises an infection modulator, an applicator, and a set of instructions.

```
L19 ANSWER 4 OF 6 USPATFULL on STN
```

AN 2003:238570 USPATFULL

TI Prostacyclin derivative containing compositions and methods of using the same for the treatment of **cancer**

IN Shorr, Robert, Edison, NJ, UNITED STATES

```
Rothblatt, Martine, Silver Spring, MD, UNITED STATES
PΙ
       US 2003166728
                          A1
                                20030904
ΑI
       US 2002-47802
                          A1
                                20020116 (10)
DT
       Utility
FS
       APPLICATION
LREP
       Allen R. Kipnes, WATOV & KIPNES, P.C., P.O. BOX 247, PRINCETON JUNCTION,
       NJ, 08550
CLMN
       Number of Claims: 28
ECL
       Exemplary Claim: 1
DRWN
       3 Drawing Page(s)
LN.CNT 1196
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       The present invention is directed to a pharmaceutical composition
       containing a cancer-treating effective amount of a class of
       prostacyclin derivatives, and a pharmaceutically acceptable carrier, and
       to kits and methods of employing the same for the treatment of
       cancer.
L19 ANSWER 5 OF 6 USPATFULL on STN
AN
       2003:159865 USPATFULL
ΤI
       Inhibitors of endothelin-1 synthesis
TN
       Corder, Roger, Harrow, UNITED KINGDOM
       Smith, Adrian P.L., London, UNITED KINGDOM
       Higenbottam, Timothy W., Sheffield, UNITED KINGDOM
       Rothblatt, Martine, Silver Spring, MD, UNITED STATES
       Vane, Sir John, London, UNITED KINGDOM
       Lees, Delphine Dominique Marthe, London, UNITED KINGDOM
PA
       United Therapeutics Corporation (non-U.S. corporation)
PΙ
       US 2003109480
                          A1
                               20030612
AΤ
       US 2002-295942
                          A1
                               20021118 (10)
RLT
       Continuation of Ser. No. US 2000-527240, filed on 17 Mar 2000, ABANDONED
PRAI
       US 1999-125000P
                           19990318 (60)
DT
       Utility
FS
       APPLICATION
       FOLEY AND LARDNER, SUITE 500, 3000 K STREET NW, WASHINGTON, DC, 20007
LREP
CLMN
       Number of Claims: 54
       Exemplary Claim: 1
ECL
       19 Drawing Page(s)
DRWN
LN.CNT 1357
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       Sequences in human preproendothelin-1 mRNA are described against which
       antisense oligonucleotides can be used to inhibit the synthesis of
       endothelin-1. This inhibition of endothelin-1 synthesis may be used to
       treat diseases where excess production of endothelin-1 is an underlying
       cause of the symptoms.
    ANSWER 6 OF 6 USPATFULL on STN
L19
       2003:158899 USPATFULL
AN
TI
       Modified prostaglandin compounds and analogs thereof, compositions
       containing the same useful for the treatment of cancer
       Shorr, Robert, Edison, NJ, UNITED STATES
IN
       Rothblatt, Martine, Silver Spring, MD, UNITED STATES
       Bentley, Michael, Huntsvillef, AL, UNITED STATES
       Zhao, Xuan, Huntsville, AL, UNITED STATES
                                                abandoned
       US 2003108512
PΙ
                          A1
                               20030612
ΑI
       US 2001-6197
                          A1
                               20011210 (10)
DT
       Utility
```

FS APPLICATION

LREP Allen R. Kipnes, WATOV & KIPNES, P.C., P.O. BOX 247, PRINCETON JUNCTION,

NJ, 08550

CLMN Number of Claims: 52

ECL Exemplary Claim: 1
DRWN 4 Drawing Page(s)

LN.CNT 1415

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

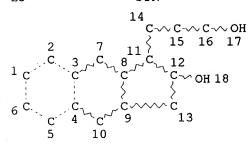
The invention is directed to a pharmaceutical composition containing a cancer-treating effective amount of a prostaglandin compound and analogs thereof having a metabolic rate slowing group attached thereto, and a pharmaceutically acceptable carrier, and methods of employing the same for the treatment of cancer.

NODE ATTRIBUTES:
CONNECT IS X3 RC AT 16
DEFAULT MLEVEL IS ATOM
DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES: RING(S) ARE ISOLATED OR EMBEDDED NUMBER OF NODES IS 18

STEREO ATTRIBUTES: NONE

L3 87 SEA FILE=REGISTRY SSS FUL L1 L5 STR



NODE ATTRIBUTES:
CONNECT IS E2 RC AT 5
CONNECT IS X3 RC AT 16
DEFAULT MLEVEL IS ATOM
DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES: RING(S) ARE ISOLATED OR EMBEDDED NUMBER OF NODES IS 18

STEREO ATTRIBUTES: NONE

L6 2 SEA FILE=REGISTRY SUB=L3 SSS FUL L5

L7 ST

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               11
                        ^ OH 18
                       13
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           21
  G2~Ak~C~G3
19
      20
               22
           0
           23
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VAR G2=O/N/S/C VAR G3=24/25 NODE ATTRIBUTES: CONNECT IS X3 RC AT CONNECT IS E2 RC AT 20 CONNECT IS E1 RC. AT 24 RC AT CONNECT IS E1 DEFAULT MLEVEL IS ATOM DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES: RING(S) ARE ISOLATED OR EMBEDDED NUMBER OF NODES IS 25

STEREO ATTRIBUTES: NONE

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L14

108 SEA FILE=EMBASE ABB=ON PLU=ON L6 OR L8
8 SEA FILE=EMBASE ABB=ON PLU=ON L14 AND (?CANCER? OR ?NEOPLAS?

OR ?CARCIN? OR ?TUMOR? OR ?TUMOUR? OR ?METAST?)

(=> d 115 ibib ab hitind 1-8)

L15 ANSWER 1 OF 8 EMBASE COPYRIGHT 2004 ELSEVIER INC. ALL RIGHTS RESERVED.

on STN

2003448000 EMBASE ACCESSION NUMBER:

New drug approvals for 2002. TITLE:

Frantz S.; Smith A. AUTHOR:

S. Frantz. s.frantz@nature.com CORPORATE SOURCE:

Nature Reviews Drug Discovery, (2003) 2/2 (95-96). SOURCE:

Refs: 1

ISSN: 1474-1776 CODEN: NRDDAG

United Kingdom COUNTRY:

DOCUMENT TYPE: Journal; Note

FILE SEGMENT: 006 Internal Medicine

> Cancer 016 Pharmacology 030

Drug Literature Index 037

English LANGUAGE:

Medical Descriptors:

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*drug approval
drug marketing
drug mechanism
food and drug administration
hypertension: DT, drug therapy
 breast metastasis: DT, drug therapy
pulmonary hypertension: DT, drug therapy
aspergillosis: DT, drug therapy
heart disease: DT, drug therapy
narcolepsy: DT, drug therapy
cataplexy: DT, drug therapy
irritable colon: DT, drug therapy
  colorectal cancer: DT, drug therapy
hepatitis B: DT, drug therapy
hypercholesterolemia: DT, drug therapy
schizophrenia: DT, drug therapy
diarrhea: DT, drug therapy
cryptosporidiosis: DT, drug therapy
giardiasis: DT, drug therapy
osteoporosis: DT, drug therapy
keratoconjunctivitis sicca: DT, drug therapy
migraine: DT, drug therapy
bacterial infection: DT, drug therapy
Alzheimer disease: DT, drug therapy
influenza: DT, drug therapy
erectile dysfunction: DT, drug therapy
osteoarthritis: DT, drug therapy
rheumatoid arthritis: DT, drug therapy
human
note
priority journal
Drug Descriptors:
*new drug: DT, drug therapy
*new drug: PD, pharmacology
olmesartan: DT, drug therapy
olmesartan: PD, pharmacology
fulvestrant: DT, drug therapy
fulvestrant: PD, pharmacology
uniprost: DT, drug therapy
uniprost: PD, pharmacology
voriconazole: DT, drug therapy
voriconazole: PD, pharmacology
dimyristoylphosphatidylcholine: DT, drug therapy
dimyristoylphosphatidylcholine: PD, pharmacology
oxybate sodium: DT, drug therapy
oxybate sodium: PD, pharmacology
tegaserod: DT, drug therapy
tegaserod: PD, pharmacology
oxaliplatin: CB, drug combination
oxaliplatin: DT, drug therapy
oxaliplatin: PD, pharmacology
fluorouracil: CB, drug combination
fluorouracil: DT, drug therapy
folinic acid: CB, drug combination
folinic acid: DT, drug therapy
adefovir dipivoxil: DT, drug therapy
adefovir dipivoxil: PD, pharmacology
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eplerenone: DT, drug therapy
eplerenone: PD, pharmacology
ezetimibe: DT, drug therapy
ezetimibe: PD, pharmacology
hypocholesterolemic agent: DT, drug therapy
hypocholesterolemic agent: PD, pharmacology
aripiprazole: DT, drug therapy
aripiprazole: PD, pharmacology
nitazoxanide: DT, drug therapy
nitazoxanide: PD, pharmacology
parathyroid hormone[1-34]: DT, drug therapy
parathyroid hormone[1-34]: PD, pharmacology
cyclosporin A: DT, drug therapy
cyclosporin A: PD, pharmacology
eletriptan: DT, drug therapy
eletriptan: PD, pharmacology
ertapenem: DT, drug therapy
ertapenem: PD, pharmacology
telmisartan: CB, drug combination
telmisartan: DT, drug therapy
telmisartan: PD, pharmacology
hydrochlorothiazide: CB, drug combination
hydrochlorothiazide: DT, drug therapy
hydrochlorothiazide: PD, pharmacology
bosentan: DT, drug therapy
bosentan: PD, pharmacology
memantine: DT, drug therapy
memantine: PD, pharmacology
oseltamivir: DT, drug therapy
oseltamivir: PD, pharmacology
tadalafil: DT, drug therapy
tadalafil: PD, pharmacology
valdecoxib: DT, drug therapy
valdecoxib: PD, pharmacology
vardenafil: DT, drug therapy
vardenafil: PD, pharmacology
unindexed drug
hepsera
inspra
alinia
micardis plus
pritor plus
ebixa
axura
levitra
(olmesartan) 144689-63-4; (fulvestrant) 129453-61-8; (uniprost)
81846-19-7; (voriconazole) 137234-62-9;
(dimyristoylphosphatidylcholine) 13699-48-4, 18194-24-6; (oxybate sodium)
502-85-2; (tegaserod) 145158-71-0, 189188-57-6; (oxaliplatin) 61825-94-3;
(fluorouracil) 51-21-8; (folinic acid) 58-05-9, 68538-85-2; (adefovir
dipivoxil) 142340-99-6; (eplerenone) 107724-20-9; (ezetimibe) 163222-33-1;
(aripiprazole) 129722-12-9; (nitazoxanide) 55981-09-4; (parathyroid
hormone[1-34]) 12583-68-5, 52232-67-4; (cyclosporin A) 59865-13-3,
63798-73-2; (eletriptan) 143322-58-1, 177834-92-3; (ertapenem)
153773-82-1, 153832-38-3, 153832-46-3; (telmisartan) 144701-48-4;
(hydrochlorothiazide) 58-93-5; (bosentan) 147536-97-8, 157212-55-0;
(memantine) 19982-08-2, 41100-52-1; (oseltamivir) 196618-13-0,
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RN

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204255-09-4, 204255-11-8; (tadalafil) 171596-29-5; (valdecoxib)
     181695-72-7; (vardenafil) 224785-90-4, 224785-91-5, 224789-15-5
     (1) Benicar; (2) Faslodex; (3) Remodulin; (4) Xyrem; (5) Zelnorm; (6)
CN
     Eloxatin; (7) Hepsera; (8) Inspra; (9) Zetia; (10) Zetia; (11) Abilify;
     (12) Abilify; (13) Alinia; (15) Forteo; (16) Relpax; (17) Invanz; (18)
     Micardis plus; (19) Pritor plus; (20) Tracleer; (21) Ebixa; (22) Axura;
     (23) Tamiflu; (24) Cialis; (25) Bextra; (26) Bextra; (27) Levitra
     (1) Sankyo; (2) Astra Zeneca; (3) United Therapeutics; (4) Orphan; (5)
CO
     Novartis; (6) Sanofi Synthelabo; (7) Gilead; (8) Searle; (9) Merck
     (Singapore); (10) Schering Plough (Singapore); (11) Otsuka; (12) Bristol
     Myers Squibb; (13) Romark; (14) Lilly; (17) Merck Sharp and Dohme; (18)
     Boehringer Ingelheim; (19) Glaxo SmithKline; (20) Actelion; (21) Lundbeck;
     (22) Merz; (23) Hoffmann La Roche; (24) Lilly ICOS; (25) Pharmacia; (26)
     Pfizer; (27) Bayer; Allergan; Alliance
                    EMBASE COPYRIGHT 2004 ELSEVIER INC. ALL RIGHTS RESERVED.
L15 ANSWER 2 OF 8
     on STN
ACCESSION NUMBER:
                    2003396573 EMBASE
                    [Pulmonary hypertension].
TITLE:
                    PULMONALE HYPERTONIE.
AUTHOR:
                    Petkov V.; Doberer D.
                    Dr. V. Petkov, Univ. Klin. fur Innere Medizin IV, Klinische
CORPORATE SOURCE:
                    Abteilung fur Pulmologie, Wahringer Gurtel 18-20, A-1090
                    Wien, Austria. Ventzislav.Petkov@univie.ac.at
                    Journal fur Hypertonie, (2003) 7/3 (7-14).
SOURCE:
                    Refs: 13
                    ISSN: 1028-2327 CODEN: JHYPFE
                    Austria
COUNTRY:
                    Journal; (Short Survey)
DOCUMENT TYPE:
                            Internal Medicine
                    006
FILE SEGMENT:
                            Chest Diseases, Thoracic Surgery and Tuberculosis
                    015
                            Cardiovascular Diseases and Cardiovascular Surgery
                    018
                    037
                            Drug Literature Index
                            Adverse Reactions Titles
                    038
LANGUAGE:
                    German
                    English; German
SUMMARY LANGUAGE:
     Pulmonary Hypertension (PH) is a haemodynamic diagnosis caused by several
     underlying diseases. In the last years tremendous progress in
     pathophysiology, diagnostics and therapy of PH was made. These new
     insights led to the first international classification of pulmonary
     hypertension (Evian 1998) and at the end of the nineties the first
     controlled trials were launched. Although further subdivision of different
     aetiologies of PH will preceed, therapeutic approaches still follow the
     haemodynamic profiles of PH. In this article we will focus on the
     precapillary forms of PH, including the prototype primary pulmonary
     hypertension, which are characterized by similar therapeutic strategies.
     Medical Descriptors:
CT
     *pulmonary hypertension: DI, diagnosis
     *pulmonary hypertension: DT, drug therapy
     *pulmonary hypertension: EP, epidemiology
     *pulmonary hypertension: ET, etiology
     *pulmonary hypertension: SI, side effect
     *pulmonary hypertension: SU, surgery
     world health organization
     disease classification
     symptomatology
```

hemodynamic parameters

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lung capillary pressure
lung artery pressure
lung vascular resistance
risk factor
disease association
drug use
drug abuse
cigarette smoking
estrogen therapy
algorithm
diagnostic test
two dimensional echocardiography
lung ventilation perfusion ratio
lung scintiscanning
computer assisted tomography
conservative treatment
lung transplantation
anticoagulant therapy
lung embolism: CO, complication
lung embolism: DT, drug therapy
lung embolism: PC, prevention
human
short survey
Drug Descriptors:
*prostacyclin derivative: AD, drug administration
*prostacyclin derivative: DO, drug dose
*prostacyclin derivative: DT, drug therapy
*prostacyclin derivative: IH, inhalational drug administration
*prostacyclin derivative: IV, intravenous drug administration
*prostacyclin derivative: PO, oral drug administration
*prostacyclin derivative: SC, subcutaneous drug administration
*calcium channel blocking agent: AD, drug administration
*calcium channel blocking agent: DT, drug therapy
*calcium channel blocking agent: PO, oral drug administration
*endothelin receptor antagonist: AD, drug administration
*endothelin receptor antagonist: DT, drug therapy
*endothelin receptor antagonist: PO, oral drug administration
*phosphodiesterase inhibitor: AD, drug administration
*phosphodiesterase inhibitor: DT, drug therapy
*phosphodiesterase inhibitor: PO, oral drug administration
*vasodilator agent: AD, drug administration
*vasodilator agent: DT, drug therapy
*vasodilator agent: PO, oral drug administration
methamphetamine: TO, drug toxicity
cocaine: TO, drug toxicity
tryptophan: AE, adverse drug reaction
aminorex: AE, adverse drug reaction
fenfluramine: AE, adverse drug reaction
  antineoplastic agent: AE, adverse drug reaction
antidepressant agent: AE, adverse drug reaction
oral contraceptive agent: AE, adverse drug reaction
oral contraceptive agent: AD, drug administration
oral contraceptive agent: PO, oral drug administration
estrogen: AE, adverse drug reaction
diltiazem: AD, drug administration
diltiazem: DT, drug therapy
diltiazem: PO, oral drug administration
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nifedipine: AD, drug administration
     nifedipine: DT, drug therapy
     nifedipine: PO, oral drug administration
     prostacyclin: AD, drug administration
     prostacyclin: DT, drug therapy
     prostacyclin: IV, intravenous drug administration
     iloprost: AD, drug administration
     iloprost: DT, drug therapy
     iloprost: IH, inhalational drug administration
     iloprost: IV, intravenous drug administration
     uniprost: AD, drug administration
     uniprost: DT, drug therapy
     uniprost: SC, subcutaneous drug administration
     beraprost: AD, drug administration
     beraprost: DT, drug therapy
     beraprost: PO, oral drug administration
     bosentan: AD, drug administration
     bosentan: DT, drug therapy
     bosentan: PO, oral drug administration
     sildenafil: AD, drug administration
     sildenafil: DT, drug therapy
     sildenafil: PO, oral drug administration
     vasoactive intestinal polypeptide: DT, drug therapy
     dihydralazine: AD, drug administration
     dihydralazine: DT, drug therapy dihydralazine: PO, oral drug administration
     urapidil: AD, drug administration
     urapidil: DT, drug therapy
     urapidil: PO, oral drug administration
     coumarin anticoagulant: AD, drug administration
     coumarin anticoagulant: DT, drug therapy
     coumarin anticoagulant: PO, oral drug administration
     digitalis: AD, drug administration
     digitalis: DT, drug therapy
     digitalis: PO, oral drug administration
     diuretic agent: AD, drug administration
     diuretic agent: DT, drug therapy
     diuretic agent: PO, oral drug administration
     dipeptidyl carboxypeptidase inhibitor: DT, drug therapy
     unindexed drug
     (methamphetamine) 28297-73-6, 51-57-0, 537-46-2, 7632-10-2; (cocaine)
     50-36-2, 53-21-4, 5937-29-1; (tryptophan) 6912-86-3, 73-22-3; (aminorex)
     13425-22-4, 2207-50-3; (fenfluramine) 404-82-0, 458-24-2; (diltiazem)
     33286-22-5, 42399-41-7; (nifedipine) 21829-25-4; (prostacyclin)
     35121-78-9, 61849-14-7; (iloprost) 78919-13-8, 82889-99-4; (uniprost)
     81846-19-7; (beraprost) 88430-50-6, 88475-69-8; (bosentan)
     147536-97-8, 157212-55-0; (sildenafil) 139755-83-2; (vasoactive intestinal
     polypeptide) 37221-79-7; (dihydralazine) 484-23-1; (urapidil) 34661-75-1;
     (digitalis) 8031-42-3, 8053-83-6
     Flolan; Ilomedin; Viagra; Tracleer
L15 ANSWER 3 OF 8 EMBASE COPYRIGHT 2004 ELSEVIER INC. ALL RIGHTS RESERVED.
     on STN
ACCESSION NUMBER:
                    2003252137 EMBASE
                    [Consensus recommendations of the working group on
TITLE:
```

RN

CN

Lung Diseases and Tuberculosis].

pulmonary arterial hypertension of the Austrian Society for

```
KONSENSUS-EMPFEHLUNGEN DER ARBEITSGRUPPE PULMONALARTERIELLE
                    HYPERTENSION DER OSTERREICHISCHEN GESELLSCHAFT FUR
                    LUNGENERKRANKUNGEN UND TUBERKULOSE.
                    Ziesche R.
AUTHOR:
                    Dr. R. Ziesche, Klinische Abteilung fur Pulmologie, Univ.
CORPORATE SOURCE:
                    Klin. fur Innere Medizin IV, Wahringer Gurtel 18-20, A-1090
                    Wien, Austria. rolf.ziesche@akh-wien.ac.at
                    Wiener Klinische Wochenschrift, (30 May 2003) 115/10
SOURCE:
                    (351 - 365).
                    Refs: 54
                    ISSN: 0043-5325 CODEN: WKWOAO
                    Austria
COUNTRY:
DOCUMENT TYPE:
                    Journal; Conference Article
                            Chest Diseases, Thoracic Surgery and Tuberculosis
FILE SEGMENT:
                    015
                    037
                            Drug Literature Index
                    038
                            Adverse Reactions Titles
LANGUAGE:
                    German
    Medical Descriptors:
     *pulmonary hypertension: DI, diagnosis
     *pulmonary hypertension: DT, drug therapy
     *pulmonary hypertension: SI, side effect
     *pulmonary hypertension: SU, surgery
     Austria
     medical society
     lung disease
     lung tuberculosis
     disease classification
     risk factor
     treatment indication
     liver toxicity: SI, side effect
     human
     conference paper
     Drug Descriptors:
     antidepressant agent: AE, adverse drug reaction
     oral contraceptive agent: AE, adverse drug reaction
       antineoplastic agent: AE, adverse drug reaction
     cocaine
     amphetamine
     aminorex: AE, adverse drug reaction
     fenfluramine: AE, adverse drug reaction
     phentermine: AE, adverse drug reaction
     tryptophan: AE, adverse drug reaction
     diltiazem: DT, drug therapy
     diltiazem: PO, oral drug administration
     nifedipine: DT, drug therapy
     nifedipine: PO, oral drug administration
     prostacyclin: DT, drug therapy
     prostacyclin: IV, intravenous drug administration
     iloprost: DT, drug therapy
     iloprost: IH, inhalational drug administration
     iloprost: IV, intravenous drug administration
     uniprost: DT, drug therapy
     uniprost: SC, subcutaneous drug administration
     beraprost: DT, drug therapy
     beraprost: PO, oral drug administration
     bosentan: AE, adverse drug reaction
     bosentan: DT, drug therapy
```

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bosentan: PO, oral drug administration
     sildenafil: DT, drug therapy
     sildenafil: PO, oral drug administration
     vasoactive intestinal polypeptide: DT, drug therapy
     vasoactive intestinal polypeptide: IH, inhalational drug administration
     dihydralazine: DT, drug therapy
     dihydralazine: PO, oral drug administration
     urapidil: DT, drug therapy
     urapidil: PO, oral drug administration
RN
     (cocaine) 50-36-2, 53-21-4, 5937-29-1; (amphetamine) 1200-47-1, 139-10-6,
     156-34-3, 2706-50-5, 300-62-9, 51-62-7, 60-13-9, 60-15-1; (aminorex)
     13425-22-4, 2207-50-3; (fenfluramine) 404-82-0, 458-24-2; (phentermine)
     1197-21-3, 122-09-8; (tryptophan) 6912-86-3, 73-22-3; (diltiazem)
     33286-22-5, 42399-41-7; (nifedipine) 21829-25-4; (prostacyclin)
     35121-78-9, 61849-14-7; (iloprost) 78919-13-8, 82889-99-4; (uniprost)
     81846-19-7; (beraprost) 88430-50-6, 88475-69-8; (bosentan)
     147536-97-8, 157212-55-0; (sildenafil) 139755-83-2; (vasoactive intestinal
     polypeptide) 37221-79-7; (dihydralazine) 484-23-1; (urapidil) 34661-75-1
     Remodulin; Flolan; Tracleer
CN
L15 ANSWER 4 OF 8 EMBASE COPYRIGHT 2004 ELSEVIER INC. ALL RIGHTS RESERVED.
     on STN
ACCESSION NUMBER:
                    2003250118 EMBASE
                    Recap of FDA product approvals - 2002.
TITLE:
SOURCE:
                    American Journal of Health-System Pharmacy, (15 Feb 2003)
                    60/4 (310+312).
                    ISSN: 1079-2082 CODEN: AHSPEK
COUNTRY:
                    United States
DOCUMENT TYPE:
                    Journal; Note
                            Public Health, Social Medicine and Epidemiology
FILE SEGMENT:
                    017
                    037
                            Drug Literature Index
                    039
                            Pharmacy
LANGUAGE:
                    English
    Medical Descriptors:
     *drug approval
     food and drug administration
     rheumatoid arthritis
     attention deficit disorder
     irritable colon
     licensing
    drug indication
     malignant neoplastic disease
    human
    note
    priority journal
     Drug Descriptors:
     *new drug
     vaccine
     orphan drug
     adalimumab
     ibritumomab tiuxetan
     fulvestrant
     oxaliplatin
     recombinant granulocyte colony stimulating factor
     rasburicase
     tegaserod
     diphtheria pertussis tetanus vaccine
```

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oxybate sodium
     nitazoxanide
     uniprost
     extraneal
     strattera
     elitek
     daptacel
     pediarix
     alinia
     orfadin
     humira
     (adalimumab) 331731-18-1; (ibritumomab tiuxetan) 206181-63-7;
RN
     (fulvestrant) 129453-61-8; (oxaliplatin) 61825-94-3; (recombinant
     granulocyte colony stimulating factor) 121181-53-1; (rasburicase)
     352311-12-7; (tegaserod) 145158-71-0, 189188-57-6; (oxybate sodium)
     502-85-2; (nitazoxanide) 55981-09-4; (uniprost) 81846-19-7
     (1) Strattera; (2) Zevalin; (3) Faslodex; (4) Eloxatin; (5) Neulasta; (6)
     Elitek; (7) Zelnorm; (8) Daptacel; (9) Pediarix; (10) Xyrem; (11) Alinia; (12) Orfadin; (13) Remodulin; (14) Extraneal; Humira
     (1) Lilly; (2) Idec; (3) Astra Zeneca; (5) Amgen; (6) Sanofi Synthelabo;
CO
     (7) Novartis; (8) Aventis Pasteur; (9) Glaxo SmithKline; (10) Orphan; (11)
     Romark; (12) Swedish Orphan; (13) United Therapeutics; (14) Baxter
L15 ANSWER 5 OF 8 EMBASE COPYRIGHT 2004 ELSEVIER INC. ALL RIGHTS RESERVED.
     on STN
                    2003080265 EMBASE
ACCESSION NUMBER:
TITLE:
                    Chronic obstructive pulmonary disease, pollution, pulmonary
                     vascular disease, transplantation, pleural disease, and
                     lung cancer in AJRCCM 2002.
                     Tobin M.J.
AUTHOR:
                     Dr. M.J. Tobin, Div. of Pulmon./Critical Care Med., Hines
CORPORATE SOURCE:
                    Veterans Affairs Hospital, Route 111N, Hines, IL 60141,
                     United States. mtobin2@lumc.edu
                     American Journal of Respiratory and Critical Care Medicine,
SOURCE:
                     (1 Feb 2003) 167/3 (356-370).
                     Refs: 98
                     ISSN: 1073-449X CODEN: AJCMED
COUNTRY:
                     United States
                     Journal; General Review
DOCUMENT TYPE:
FILE SEGMENT:
                     015
                             Chest Diseases, Thoracic Surgery and Tuberculosis
                     016
                     017
                             Public Health, Social Medicine and Epidemiology
                     037
                             Drug Literature Index
                             Adverse Reactions Titles
                     038
                     English
LANGUAGE:
     Medical Descriptors:
     *chronic obstructive lung disease: DT, drug therapy
     *chronic obstructive lung disease: EP, epidemiology
     *chronic obstructive lung disease: ET, etiology
     *chronic obstructive lung disease: SU, surgery
     *chronic obstructive lung disease: TH, therapy
     *lung transplantation
     *pulmonary hypertension: DT, drug therapy
     *pulmonary hypertension: ET, etiology
     *lung embolism: DI, diagnosis
     *lung embolism: ET, etiology
     *pleura disease: DI, diagnosis
```

```
*pleura disease: ET, etiology
*pleura disease: SU, surgery
*air pollution
pathogenesis
genetic polymorphism
risk factor
alpha 1 antitrypsin deficiency: DT, drug therapy
alpha 1 antitrypsin deficiency: ET, etiology
mortality
oxygen therapy
smoking
pneumonia: ET, etiology
pathophysiology
breathing
breathing muscle
cardiovascular disease: SI, side effect
supraventricular tachycardia: SI, side effect
drug efficacy
drug megadose
corticosteroid therapy
corticosteroid induced osteoporosis: SI, side effect
vertebra fracture: SI, side effect
hip fracture: SI, side effect
proteinase inhibition
headache: SI, side effect
hyperlipidemia: SI, side effect
myalgia: SI, side effect
treatment outcome
asthma: DT, drug therapy
sickle cell anemia: ET, etiology
treatment indication
patient selection
bronchiolitis: ET, etiology
  lung cancer: DI, diagnosis
  lung cancer: EP, epidemiology
human
nonhuman
clinical trial
review
priority journal
Drug Descriptors:
technetium 99m
muscarinic receptor blocking agent: AE, adverse drug reaction
muscarinic receptor blocking agent: DT, drug therapy
ipratropium bromide: AE, adverse drug reaction
ipratropium bromide: DT, drug therapy
placebo
theophylline: CT, clinical trial
theophylline: DO, drug dose
theophylline: DT, drug therapy
theophylline: PD, pharmacology
glucocorticoid: AE, adverse drug reaction
glucocorticoid: CT, clinical trial
glucocorticoid: CM, drug comparison
glucocorticoid: DO, drug dose
glucocorticoid: DT, drug therapy
glucocorticoid: IH, inhalational drug administration
```

```
glucocorticoid: PO, oral drug administration
budesonide: AE, adverse drug reaction
budesonide: CT, clinical trial
budesonide: DT, drug therapy
budesonide: IH, inhalational drug administration
prednisolone: AE, adverse drug reaction
prednisolone: CT, clinical trial
prednisolone: DT, drug therapy
prednisolone: PO, oral drug administration
fluticasone propionate: CT, clinical trial
fluticasone propionate: CB, drug combination
fluticasone propionate: CM, drug comparison
fluticasone propionate: DO, drug dose
fluticasone propionate: DT, drug therapy
fluticasone propionate: IH, inhalational drug administration
salmeterol: CT, clinical trial
salmeterol: CB, drug combination
salmeterol: CM, drug comparison
salmeterol: DO, drug dose
salmeterol: DT, drug therapy
salmeterol: IH, inhalational drug administration
proteinase inhibitor: AE, adverse drug reaction
proteinase inhibitor: CT, clinical trial
proteinase inhibitor: CR, drug concentration
proteinase inhibitor: DT, drug therapy
proteinase inhibitor: PD, pharmacology
proteinase inhibitor: PO, oral drug administration
retinoic acid: AE, adverse drug reaction
retinoic acid: CT, clinical trial
retinoic acid: CR, drug concentration
retinoic acid: DT, drug therapy
retinoic acid: PD, pharmacology
ono 6818: PD, pharmacology
ono 6818: PO, oral drug administration
zd 0892: PD, pharmacology
alpha 1 antitrypsin: DT, drug therapy
ascorbic acid: DT, drug therapy
alpha tocopherol: DT, drug therapy
norfloxacin: PD, pharmacology
prostacyclin: DT, drug therapy
nitric oxide: DT, drug therapy
uniprost: CT, clinical trial
uniprost: DT, drug therapy
uniprost: SC, subcutaneous drug administration
monocrotaline: DT, drug therapy
monocrotaline: PD, pharmacology
simvastatin: DT, drug therapy
simvastatin: PD, pharmacology
3 hydroxy 3 methylglutaryl coenzyme A: DT, drug therapy
3 hydroxy 3 methylglutaryl coenzyme A: PD, pharmacology
sildenafil: DT, drug therapy
sildenafil: PD, pharmacology
phosphodiesterase inhibitor: DT, drug therapy
phosphodiesterase inhibitor: PD, pharmacology
D dimer: EC, endogenous compound
cyclosporin: PD, pharmacology
unclassified drug
```

RN (technetium 99m) 14133-76-7; (ipratropium bromide) 22254-24-6; (theophylline) 58-55-9, 5967-84-0, 8055-07-0, 8061-56-1, 99007-19-9; (budesonide) 51333-22-3; (prednisolone) 50-24-8; (fluticasone propionate) 80474-14-2; (salmeterol) 89365-50-4; (proteinase inhibitor) 37205-61-1; (retinoic acid) 302-79-4; (alpha 1 antitrypsin) 9041-92-3; (ascorbic acid) 134-03-2, 15421-15-5, 50-81-7; (alpha tocopherol) 1406-18-4, 1406-70-8, 52225-20-4, 58-95-7, 59-02-9; (norfloxacin) 70458-96-7; (prostacyclin) 35121-78-9, 61849-14-7; (nitric oxide) 10102-43-9; (uniprost) 81846-19-7; (monocrotaline) 315-22-0, 8051-27-2; (simvastatin) 79902-63-9; (3 hydroxy 3 methylglutaryl coenzyme A) 1553-55-5; (sildenafil) 139755-83-2; (cyclosporin) 79217-60-0

CN Ono 6818; Zd 0892; Prolastin

L15 ANSWER 6 OF 8 EMBASE COPYRIGHT 2004 ELSEVIER INC. ALL RIGHTS RESERVED.

on STN

ACCESSION NUMBER: 2003072829 EMBASE

TITLE: Gateways to clinical trials.

AUTHOR: Bayes M.; Rabasseda X.; Prous J.R.

CORPORATE SOURCE: M. Bayes, Prous Science, P.O. Box 540, 08080 Barcelona,

Spain. mbayes@prous.com

SOURCE: Methods and Findings in Experimental and Clinical

Pharmacology, (2002) 24/10 (703-729). after 1/10/02

Refs: 180

ISSN: 0379-0355 CODEN: MFEPDX

COUNTRY: Spain

DOCUMENT TYPE: Journal; General Review

FILE SEGMENT: 018 Cardiovascular Diseases and Cardiovascular Surgery

037 Drug Literature Index 038 Adverse Reactions Titles

048 Gastroenterology

LANGUAGE: English SUMMARY LANGUAGE: English

Gateways to Clinical Trials is a guide to the most recent clinical trials in current literature and congresses. The data in the following tables has been retrieved from the Clinical Studies knowledge area of Prous Science Integrity, the drug discovery and development portal, http://integrity.prous.com. This issue focuses on the following selection of drugs: Abacavir sulfate, adalimumab, AERx morphine sulphate, alefacept, alemtuzumab, alendronic acid sodium salt, alicaforsen sodium, almotriptan, amprenavir, aripiprazole, atenolol, atorvastatin calcium; BSYX-A110; Cantuzumab mertansine, capravirine, CDP-571, CDP-870, celecoxib; Delavirdine mesilate, docetaxel, dofetilide, donepezil hydrochloride, duloxetine hydrochloride, dutasteride, dydrogesterone; Efavirenz, emtricitabine, enjuvia, enteryx, epristeride, erlotinib hydrochloride, escitalopram oxalate, etanercept, etonogestrel, etoricoxib; Fesoterodine, finasteride, flt3ligand; Galantamine hydrobromide, gemtuzumab ozogamicin, genistein, gepirone hydrochloride; Indinavir sulfate, infliximab; Lamivudine, lamivudine/zidovudine/abacavir sulfate, leteprinim potassium, levetiracetam, liposomal doxorubicin, lopinavir, lopinavir/ritonavir, losartan potassium; MCC-465, MRA; Nebivolol, nesiritide, nevirapine; Olanzapine, OROS(R)-Methylphenidate hydrochloride; Peginterferon alfa-2a, peginterferon alfa-2b, Pimecrolimus, polyethylene glycol 3350, pramlintide acetate, pregabalin, PRO-2000; Risedronate sodium, risperidone, ritonavir, rituximab, rivastigmine tartrate, rofecoxib, rosuvastatin calcium; Saquinavir mesilate, Stavudine; Tacrolimus, tadalafil, tamsulosin hydrochloride, telmisartan, tomoxetine hydrochloride, treprostinil sodium, trimegestone, trimetrexate; Valdecoxib, venlafaxine hydrochloride;

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Zoledronic acid monohydrate. .COPYRGT. 2002 Prous Science. All rights
     reserved.
CT
    Medical Descriptors:
     *drug research
     medical literature
     cardiovascular disease: DT, drug therapy
     dose response
     side effect: SI, side effect
     gastrointestinal disease: DT, drug therapy
     diarrhea: SI, side effect
     virus infection: DT, drug therapy
     metabolic disorder: DT, drug therapy
     nutritional disorder: DT, drug therapy
     musculoskeletal disease: DT, drug therapy
     connective tissue disease
     bone disease: DT, drug therapy
       neoplasm: DT, drug therapy
     neurologic disease: DT, drug therapy
     mental disease: DT, drug therapy
     kidney disease: DT, drug therapy
     genital system disease: DT, drug therapy
     thrombophlebitis: SI, side effect
     breast disease: DT, drug therapy
     skin disease: DT, drug therapy
    human
     clinical trial
    meta analysis
     review
    Drug Descriptors:
    nesiritide: CT, clinical trial
    nesiritide: CB, drug combination
nesiritide: CM, drug comparison
nesiritide: DT, drug therapy
    nesiritide: IV, intravenous drug administration
    glyceryl trinitrate: CT, clinical trial
    glyceryl trinitrate: CM, drug comparison
    glyceryl trinitrate: DT, drug therapy
    glyceryl trinitrate: IV, intravenous drug administration
    milrinone: CT, clinical trial
    milrinone: CB, drug combination
    milrinone: DT, drug therapy
    dofetilide: CT, clinical trial
    dofetilide: DT, drug therapy
    losartan: CT, clinical trial
    losartan: DT, drug therapy
    atenolol: CT, clinical trial
    atenolol: DT, drug therapy
    ascorbic acid: CT, clinical trial
    ascorbic acid: CM, drug comparison
    ascorbic acid: DT, drug therapy
    ascorbic acid: PO, oral drug administration
    atorvastatin: CT, clinical trial
    atorvastatin: CM, drug comparison
    atorvastatin: DT, drug therapy
    atorvastatin: PO, oral drug administration
    enalapril: CT, clinical trial
    enalapril: CM, drug comparison
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enalapril: DT, drug therapy
telmisartan: CT, clinical trial
telmisartan: CM, drug comparison
telmisartan: DT, drug therapy
nebivolol: CT, clinical trial
nebivolol: DT, drug therapy
uniprost: CT, clinical trial
uniprost: DO, drug dose
uniprost: DT, drug therapy
rosuvastatin: AE, adverse drug reaction
rosuvastatin: CT, clinical trial rosuvastatin: DT, drug therapy
macrogol: AE, adverse drug reaction
macrogol: CT, clinical trial
macrogol: DO, drug dose
macrogol: DT, drug therapy
alicaforsen: CT, clinical trial
alicaforsen: DT, drug therapy
alicaforsen: IV, intravenous drug administration
  tumor necrosis factor alpha antibody: AE, adverse drug reaction
  tumor necrosis factor alpha antibody: CT, clinical trial
  tumor necrosis factor alpha antibody: DT, drug therapy
  tumor necrosis factor alpha antibody: IV, intravenous drug
administration
etoricoxib: CT, clinical trial
etoricoxib: CM, drug comparison
etoricoxib: DT, drug therapy
etoricoxib: PO, oral drug administration
rofecoxib: CT, clinical trial
rofecoxib: CM, drug comparison
rofecoxib: DT, drug therapy
diclofenac: CT, clinical trial
diclofenac: CM, drug comparison
diclofenac: DT, drug therapy
infliximab: CT, clinical trial
infliximab: CB, drug combination
infliximab: DT, drug therapy
infliximab: IV, intravenous drug administration
prednisone: CT, clinical trial
prednisone: CB, drug combination
prednisone: DT, drug therapy
peginterferon alpha2a: CT, clinical trial
peginterferon alpha2a: CB, drug combination
peginterferon alpha2a: DT, drug therapy
peginterferon alpha2a: SC, subcutaneous drug administration
ribavirin: CT, clinical trial
ribavirin: CB, drug combination
ribavirin: IT, drug interaction
peginterferon alpha2b: CT, clinical trial
peginterferon alpha2b: CB, drug combination
peginterferon alpha2b: DT, drug therapy
lamivudine: CT, clinical trial lamivudine: CB, drug combination lamivudine: DT, drug therapy
lamivudine: PO, oral drug administration
abacavir: CT, clinical trial
abacavir: CB, drug combination
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abacavir: DT, drug therapy
     zidovudine: CT, clinical trial
     zidovudine: CB, drug combination
     zidovudine: DT, drug therapy
     stavudine: CT, clinical trial
     stavudine: CB, drug combination
     stavudine: DT, drug therapy
     delavirdine: CT, clinical trial
     delavirdine: CB, drug combination
     delavirdine: DT, drug therapy
     unindexed drug
RN
     (nesiritide) 124584-08-3, 189032-40-4; (glyceryl trinitrate) 55-63-0;
     (milrinone) 78415-72-2; (dofetilide) 115256-11-6; (losartan) 114798-26-4;
     (atenolol) 29122-68-7; (ascorbic acid) 134-03-2, 15421-15-5, 50-81-7;
     (atorvastatin) 134523-00-5, 134523-03-8; (enalapril) 75847-73-3;
     (telmisartan) 144701-48-4; (nebivolol) 99200-09-6; (uniprost)
     81846-19-7; (rosuvastatin) 147098-18-8, 147098-20-2; (macrogol)
     25322-68-3; (alicaforsen) 142442-63-5, 185229-68-9, 331257-52-4;
     (etoricoxib) 202409-33-4, 202409-40-3; (rofecoxib) 162011-90-7,
     186912-82-3; (diclofenac) 15307-79-6, 15307-86-5; (infliximab)
     170277-31-3; (prednisone) 53-03-2; (peginterferon alpha2a) 198153-51-4;
     (ribavirin) 36791-04-5; (peginterferon alpha2b) 215647-85-1; (lamivudine)
     134678-17-4, 134680-32-3; (abacavir) 136470-78-5, 188062-50-2;
     (zidovudine) 30516-87-1; (stavudine) 3056-17-5; (delavirdine) 136817-59-9
CN
     Cdp 571
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     on STN
ACCESSION NUMBER:
                    2002412976 EMBASE
TITLE:
                    News focus.
SOURCE:
                    Current Drug Discovery, (1 Nov 2002) -/NOV. (11).
                    ISSN: 1472-7463 CODEN: CDDUAI
COUNTRY:
                    United Kingdom
DOCUMENT TYPE:
                    Journal; Note
FILE SEGMENT:
                    003
                            Endocrinology
                            Microbiology
                    004
                    016
                            Cancer
                    018
                            Cardiovascular Diseases and Cardiovascular Surgery
                            Pharmacology
                    030
                    037
                            Drug Literature Index
LANGUAGE:
                    English
    Medical Descriptors:
     non insulin dependent diabetes mellitus: DT, drug therapy
     glucose blood level
    premature labor: DT, drug therapy
     ovary insufficiency
     fertilization in vitro
     growth hormone deficiency: DT, drug therapy
      solid tumor: DT, drug therapy
      antineoplastic activity
     aspergillosis: DT, drug therapy
    bacterial infection: DT, drug therapy
    drug potency
    drug structure
    antibacterial activity
    pulmonary hypertension: DT, drug therapy
    human
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controlled study
Drug Descriptors:
*protein tyrosine phosphatase inhibitor: DT, drug therapy
*protein tyrosine phosphatase inhibitor: PD, pharmacology
*protein tyrosine phosphatase inhibitor: PO, oral drug administration
  *antineoplastic agent: DT, drug therapy
  *antineoplastic agent: PD, pharmacology
*antiinfective agent: AN, drug analysis
*antiinfective agent: CM, drug comparison
*antiinfective agent: DT, drug therapy
*antiinfective agent: PD, pharmacology
*antiinfective agent: IV, intravenous drug administration
as 602305: DT, drug therapy
as 602305: PO, oral drug administration
oxytocin antagonist: DT, drug therapy
oxytocin antagonist: PO, oral drug administration
recombinant follitropin: PD, pharmacology
sermorelin: DT, drug therapy
abt 100: DT, drug therapy
abt 100: PD, pharmacology
abt 839
protein farnesyltransferase inhibitor: DT, drug therapy
protein farnesyltransferase inhibitor: PD, pharmacology
abt 567: PD, pharmacology
angiogenesis inhibitor: PD, pharmacology
n [[5 [(2 amino 3 mercaptopropyl)amino][1,1' biphenyl] 2
yl]carbonyl]methionine
bms 379224: DT, drug therapy
bms 379224: IV, intravenous drug administration
antifungal agent: DT, drug therapy
antifungal agent: IV, intravenous drug administration
ravuconazole
chorismate synthase inhibitor: DT, drug therapy
chorismate synthase inhibitor: PD, pharmacology
phosphopantethiene adenyltransferase inhibitor: DT, drug therapy
phosphopantethiene adenyltransferase inhibitor: PD, pharmacology
ptx 110130: CM, drug comparison
ptx 110130: PD, pharmacology
ptx 008313: AN, drug analysis
ptx 008313: CM, drug comparison
ptx 008313: PD, pharmacology
a 00000764: AN, drug analysis
a 00000764: PD, pharmacology
a 00026158: PD, pharmacology
a 00000762: PD, pharmacology
ar 328: PD, pharmacology
antibiotic agent: AN, drug analysis
antibiotic agent: CM, drug comparison
antibiotic agent: DT, drug therapy
antibiotic agent: PD, pharmacology
uniprost: DT, drug therapy
uniprost: SC, subcutaneous drug administration
gepirone: PD, pharmacology
mirtazapine
antidepressant agent: PD, pharmacology
unindexed drug
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unclassified drug
     (sermorelin) 86168-78-7; (n [[5 [(2 amino 3 mercaptopropyl)amino][1,1'
RN
     biphenyl] 2 yl]carbonyl]methionine) 170006-72-1; (ravuconazole)
     182760-06-1; (uniprost) 81846-19-7; (gepirone) 83928-66-9,
     83928-76-1; (mirtazapine) 61337-67-5
     (1) As 602305; (2) Abt 100; (3) Abt 567; (4) Abt 839; (5) Fti 276; (6) Bms
CN
     379224; (7) Ptx 110130; (8) Ptx 008313; (9) A 00000764; (10) A 00000762;
     (11) A 00026158; (12) Ar 328; (13) Ariza; (14) Remeron
     (1) Serono; (4) Abbott; (5) University of Pittsburgh; (6) Bristol Myers
CO
     Squibb; (8) PanTherix; (11) Arrow; (12) Arpida; (14) Organon
L15 ANSWER 8 OF 8 EMBASE COPYRIGHT 2004 ELSEVIER INC. ALL RIGHTS RESERVED.
     on STN
ACCESSION NUMBER:
                    2002123171 EMBASE
                    News focus.
TITLE:
                    Current Drug Discovery, (2002) -/MAR. (15-18).
SOURCE:
                    ISSN: 1472-7463 CODEN: CDDUAI
                    United Kingdom
COUNTRY:
                    Journal; Article
DOCUMENT TYPE:
                    016
                            Cancer
FILE SEGMENT:
                             Public Health, Social Medicine and Epidemiology
                    017
                            Pharmacology
                    030
                            Drug Literature Index
                    037
                    039
                             Pharmacy
LANGUAGE:
                    English
     Medical Descriptors:
     multidrug resistance
     degenerative disease: DT, drug therapy
     diabetes mellitus: DT, drug therapy .
     graft versus host reaction: DT, drug therapy
     bacterial infection: DT, drug therapy
     Human immunodeficiency virus infection: DT, drug therapy
       basal cell carcinoma: DT, drug therapy
     actinic keratosis: DT, drug therapy
     pulmonary hypertension: DT, drug therapy
     melanoma: DT, drug therapy
     cystic fibrosis: DT, drug therapy
       colorectal cancer: DT, drug therapy
     heart failure: DT, drug therapy
     obesity: DT, drug therapy
     erectile dysfunction: DT, drug therapy
     female sexual dysfunction: DT, drug therapy
     hip fracture: DT, drug therapy
     human
     nonhuman
     clinical trial
     controlled study
     article
     Drug Descriptors:
     *new drug: CT, clinical trial
*new drug: AN, drug analysis
     *new drug: DT, drug therapy
     *new drug: PR, pharmaceutics
     *new drug: PD, pharmacology
     *new drug: NA, intranasal drug administration
     *new drug: PO, oral drug administration
     protein inhibitor: CT, clinical trial
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protein inhibitor: DT, drug therapy
ont 093: CT, clinical trial
ont 093: DT, drug therapy
glutamate receptor antagonist: DV, drug development
glutamate receptor antagonist: DT, drug therapy
ro 68 0921: DV, drug development
ro 68 0921: DT, drug therapy
ro 64 5229: DV, drug development
ro 64 5229: DT, drug therapy
metalloproteinase inhibitor: DV, drug development
metalloproteinase inhibitor: DT, drug therapy
kb r7785: DV, drug development
kb r7785: DT, drug therapy
antiinfective agent: AN, drug analysis
antiinfective agent: DV, drug development
antiinfective agent: PD, pharmacology
pyrrole derivative: AN, drug analysis
pyrrole derivative: DV, drug development
pyrrole derivative: PD, pharmacology
tenofovir disoproxil: DT, drug therapy
  antineoplastic agent: DT, drug therapy
  antineoplastic agent: PR, pharmaceutics
metvix pdt: DT, drug therapy
metvix pdt: PR, pharmaceutics
uniprost: CT, clinical trial
uniprost: DT, drug therapy
doxycycline: PO, oral drug administration
bacterial vaccine: DT, drug therapy
aerugen: DT, drug therapy
  cancer vaccine: CT, clinical trial
  cancer vaccine: DT, drug therapy
oncophage: DT, drug therapy
theratope: CT, clinical trial
theratope: DT, drug therapy
angiogenesis inhibitor: CT, clinical trial
angiogenesis inhibitor: DT, drug therapy
semaxinib: CT, clinical trial
semaxinib: DT, drug therapy
endothelin receptor antagonist: CT, clinical trial
endothelin receptor antagonist: DT, drug therapy
bosentan: CT, clinical trial
bosentan: DT, drug therapy
antiobesity agent: CT, clinical trial
antiobesity agent: DT, drug therapy
aod 9604: CT, clinical trial
aod 9604: DT, drug therapy
apomorphine: DT, drug therapy
apomorphine: NA, intranasal drug administration
growth hormone releasing factor: DT, drug therapy
th 9507: DT, drug therapy
unindexed drug
unclassified drug
pennsaid
viread
(tenofovir disoproxil) 202138-50-9; (uniprost) 81846-19-7;
 (doxycycline) 10592-13-9, 17086-28-1, 564-25-0; (apomorphine) 314-19-2,
58-00-4; (growth hormone releasing factor) 83930-13-6, 9034-39-3
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- (1) Ont 093; (2) Ro 68 0921; (3) Ro 64 5229; (4) Kb r7785; (5) Metvix pdt; CN(6) Remodulin; (7) Oncophage; (8) Periostat; (9) Pennsaid; (10) Aerugen;
 - (11) Theratope; (12) Semaxinib; (13) Tracleer; Viread
- (1) Ontogen; (3) Hoffmann La Roche; (4) Organon; (5) PhotoCure; (6) United Therapeutics; (7) Antigenics; (8) Collagenex; (9) Dimethaid; (10) Berna; (11) Biomira; (12) Pharmacia; (13) Genentech; Genelabs; Nastech CO